

LMT/pdd
04/02/01

Date: <u>4/5/01</u> EXPRESS MAIL LABEL NO. <u>EL5522 85593US</u>

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Leonard P. Guarente, Nicanor Austriaco, Jr., James J. Claus, Francesca Cole and Brian Kennedy

Title: GENES DETERMINING CELLULAR SENESCENCE IN YEAST

TRANSMITTAL OF SEQUENCE LISTING IN COMPUTER READABLE FORM
IN COMPLIANCE WITH 37 C.F.R. §§1.821(e) AND (f)

Box Sequence
Assistant Commissioner for Patents
Washington, D.C. 20231

Sir:

Transmitted herewith is a copy of the "Sequence Listing" in computer readable form as required by 37 C.F.R. §1.821(e). As required by 37 C.F.R. §1.821(f), Applicants' Attorney hereby states that the content of the "Sequence Listing" in paper form and of the computer readable form of the "Sequence Listing" are the same.

Respectfully submitted,

HAMILTON, BROOK, SMITH & REYNOLDS, P.C.

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Date: 4/5/01

00501491-005

SEQUENCE LISTING

<110> Guarente, Leonard P.
Austriaco Jr., Nicanor
Claus, James J.
Cole, Francesca
Kennedy, Brian

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YEAST

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<151> 1995-02-28

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 Glu Gln Pro Ser Asp Gly Arg Ser Val Gly Gly Leu Tyr Cys Lys Asn
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 Gly Lys Leu Tyr Arg Ser Asn Thr Asp Thr Asn Ser Leu Cys Val Glu
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 aac atg ctg acg gaa gag ctc tat ggg aac aca ttt cag ctt tac aag 850
 Ala Ile Val Glu Tyr Ala Tyr Asn Asp Lys Ala Ile Leu Glu Gln Arg 125
 115 120
 tca gca gat cac cga act ctg gac aaa gtg tta gag gta cag cca gaa 898
 Asn Met Leu Thr Glu Glu Leu Tyr Gly Asn Thr Phe Gln Leu Tyr Lys 140
 130 135
 aaa tta gaa ctt att atg gat gaa atg aaa cag att cta act cca atg 946
 Ser Ala Asp His Arg Thr Leu Asp Lys Val Leu Glu Val Gln Pro Glu 160
 145 150 155

gcc caa aag gaa gct gtg att aag cac tca ttg gtg cat aaa gta ttc 994
 Lys Leu Glu Leu Ile Met Asp Glu Met Lys Gln Ile Leu Thr Pro Met 175
 165

ttg gac ttt ttt acc tat gca ccc ccc aaa ctc aga tca gaa atg att 1042
 Ala Gln Lys Glu Ala Val Ile Lys His Ser Leu Val His Lys Val Phe 190
 180

gaa gcc atc cgc gaa gcg gtg gtc tac ctg gca cac aca cac gat ggc 1090
 Leu Asp Phe Phe Thr Tyr Ala Pro Lys Leu Arg Ser Glu Met Ile 205
 195

gcc aga gtg gcc atg cac tgc ctg tgg cat ggc acg ccc aag gac agg 1138
 Glu Ala Ile Arg Glu Ala Val Val Tyr Leu Ala His Thr His Asp Gly 220
 210

aaa gtg att gtg aaa aca atg aag act tat gtt gaa aag gtg gct aat 1186
 Ala Arg Val Ala Met His Cys Leu Trp His Gly Thr Pro Lys Asp Arg 240
 225

ggc caa tac tcc cat ttg gtt tta ctg gcg gca ttt gat tgt att gat 1234
 Lys Val Ile Val Lys Thr Met Lys Thr Tyr Val Glu Lys Val Ala Asn 255
 245

gat act aag ctt gtg aag cag ata atc ata tca gaa att atc agt tca 1282
 Gly Gln Tyr Ser His Leu Val Leu Leu Ala Ala Phe Asp Cys Ile Asp 270
 260

ttg cct agc ata gta aat gac aaa tat gga agg aag gtc cta ttg tac 1330
 Asp Thr Lys Leu Val Lys Gln Ile Ile Ser Glu Ile Ile Ser Ser 285
 275

tta cta agc ccc aga gat cct gca cat aca gta cga gaa atc att gaa 1378
 Leu Pro Ser Ile Val Asn Asp Lys Tyr Gly Arg Lys Val Leu Leu Tyr 300
 290

gtt ctg caa aaa gga gat gga aat gca cac agt aag aaa gat aca gag 1426
 Leu Leu Ser Pro Arg Asp Pro Ala His Thr Val Arg Glu Ile Ile Glu 320
 305

gtc cgc aga cgg gag ctc cta gaa tcc att tct cca gct ttg tta agc 1474
 Val Leu Gln Lys Gly Asp Gly Asn Ala His Ser Lys Lys Asp Thr Glu 335
 325

tac ctg caa gaa cac gcc caa gaa gtg gtg cta gat aag tct gcg tgt 1522
 Val Arg Arg Arg Glu Leu Leu Glu Ser Ile Ser Pro Ala Leu Leu Ser 350
 340

gtg ttg gtg tct gac att ctg gga tct gcc act gga gac gtt cag cct 1570
 Tyr Leu Gln Glu His Ala Gln Glu Val Val Leu Asp Lys Ser Ala Cys 365
 355

acc atg aat gcc atc gcc agc ttg gca gca aca gga ctg cat cct ggt 1618
 Val Leu Val Ser Asp Ile Leu Gly Ser Ala Thr Gly Asp Val Gln Pro 380
 370

ggc aag gac gga gag ctt cac att gca gaa cat cct gca gga cat cta 1666
 Thr Met Asn Ala Ile Ala Ser Leu Ala Ala Thr Gly Leu His Pro Gly

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<210> 10
<211> 508
<212> PRT
<213> Homo sapiens
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 180 185 190
 Leu Asp Phe Phe Thr Tyr Ala Pro Pro Lys Leu Arg Ser Glu Met Ile
 195 200 205
 Glu Ala Ile Arg Glu Ala Val Val Tyr Leu Ala His Thr His Asp Gly
 210 215 220
 Ala Arg Val Ala Met His Cys Leu Trp His Gly Thr Pro Lys Asp Arg
 225 230 235 240
 Lys Val Ile Val Lys Thr Met Lys Thr Tyr Val Glu Lys Val Ala Asn
 245 250 255
 Gly Gln Tyr Ser His Leu Val Leu Leu Ala Ala Phe Asp Cys Ile Asp
 260 265 270
 Asp Thr Lys Leu Val Lys Gln Ile Ile Ile Ser Glu Ile Ile Ser Ser
 275 280 285
 Leu Pro Ser Ile Val Asn Asp Lys Tyr Gly Arg Lys Val Leu Leu Tyr
 290 295 300
 Leu Leu Ser Pro Arg Asp Pro Ala His Thr Val Arg Glu Ile Ile Glu
 305 310 315 320
 Val Leu Gln Lys Gly Asp Gly Asn Ala His Ser Lys Lys Asp Thr Glu
 325 330 335
 Val Arg Arg Arg Glu Leu Leu Glu Ser Ile Ser Pro Ala Leu Leu Ser
 340 345 350
 Tyr Leu Gln Glu His Ala Gln Glu Val Val Leu Asp Lys Ser Ala Cys
 355 360 365
 Val Leu Val Ser Asp Ile Leu Gly Ser Ala Thr Gly Asp Val Gln Pro
 370 375 380
 Thr Met Asn Ala Ile Ala Ser Leu Ala Ala Thr Gly Leu His Pro Gly
 385 390 395 400
 Gly Lys Asp Gly Glu Leu His Ile Ala Glu His Pro Ala Gly His Leu
 405 410 415
 Val Leu Lys Trp Leu Ile Glu Gln Asp Lys Lys Met Lys Glu Asn Gly
 420 425 430
 Arg Glu Gly Cys Phe Ala Lys Thr Leu Val Glu His Val Gly Met Lys
 435 440 445
 Asn Leu Lys Ser Trp Ala Ser Val Asn Arg Gly Ala Ile Ile Leu Ser
 450 455 460
 Ser Leu Leu Gln Ser Cys Asp Leu Glu Val Ala Asn Lys Val Lys Ala
 465 470 475 480
 Ala Leu Lys Ser Leu Ile Pro Thr Leu Glu Lys Thr Lys Ser Thr Ser
 485 490 495
 Lys Gly Ile Glu Ile Leu Leu Glu Lys Leu Ser Thr
 500 505

<210> 11
 <211> 2457
 <212> DNA
 <213> *Saccharomyces cerevisiae*

<220>
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1

5

25

40

55

70

85

AS

As:

610

16

Ly

17

<400> 12
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 Phe Ser Ala Pro Ala Pro Ala Pro Ala Asp Ser His His Glu Asp His
 20 25 30
 His Lys Asp Glu Lys Pro Ala Val Val Thr Val Thr Gln Tyr Ile Asp
 35 40 45
 Ser Asn Ala Ala Thr Ser Thr Val Glu Ser Ala Ala Thr Thr Thr Thr
 50 55 60
 Leu Ser Ser Ser Glu Lys Asp Thr Ser Glu Gln Lys Arg Asp Gly Gly
 65 70 75 80
 Phe Gln Asp Gly Thr Val Lys Cys Ser Asp Phe Pro Ser Val Asn Gly
 85 90 95
 Ile Val Ser Leu Asp Trp Leu Gly Phe Gly Gly Trp Ala Ser Val Met
 100 105 110
 Asp Met Asp Ala Asn Thr Ser Ser Glu Cys Lys Asp Gly Tyr Tyr Cys
 115 120 125
 Ser Tyr Ala Cys Glu Pro Gly Met Ser Lys Thr Gln Trp Pro Ser Asp
 130 135 140
 Gln Pro Ser Asp Gly Lys Ser Val Gly Gly Leu Tyr Cys Lys Asn Gly
 145 150 155 160
 Tyr Leu Tyr Arg Thr Asn Thr Asp Thr Ser Asp Leu Cys Ser Thr Asp
 165 170 175
 Glu Thr Ser Ala Lys Ala Ile Asn Lys Lys Ser Asp Ser Ile Ala Leu
 180 185 190
 Cys Arg Thr Asp Tyr Pro Gly Ser Glu Asn Met Val Ile Pro Thr Val
 195 200 205
 Val Asp Gly Gly Asp Ser Gln Pro Ile Ser Val Val Asp Glu Asp Thr
 210 215 220
 Tyr Tyr Gln Trp Gln Gly Lys Lys Thr Ser Ala Gln Tyr Tyr Ile Asn
 225 230 235 240
 Asn Ala Gly Val Ser Ala Glu Asp Gly Cys Ile Trp Gly Thr Ser Gly
 245 250 255
 Ser Asp Val Gly Asn Trp Ala Pro Leu Val Leu Gly Ala Gly Ser Thr
 260 265 270
 Asn Gly Glu Thr Tyr Leu Ser Leu Ile Pro Asn Pro Asn Ser Asn Gln
 275 280 285
 Ala Ala Asn Phe Asn Val Lys Ile Val Ala Ser Asp Gly Ala Asn Val
 290 295 300
 Gln Gly Ser Cys Ala Tyr Glu Asp Gly Ser Phe Thr Gly Asp Gly Ser
 305 310 315 320
 Asp Gly Cys Thr Val Ser Val Leu Ser Gly Ser Ala Glu Phe Val Phe
 325 330 335
 Tyr

<210> 13
 <211> 2150
 <212> DNA
 <213> *Saccharomyces cerevisiae*

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 <223> SAG1

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 tgccatgaat accaaggctt gctcagcttc tgcagcagtt caaccctttc caataccgcc 180

aatgcgtcct caaaacgtta gtttagtcgt gctcaaccgc tatttttggt tttatcttcg 240
 tttctttctc ctgaacgaca ttcgtcacga aaattgcggc ggaaaatttc ctgatgcgga 300
 cactttttcc cgatccggac atgccttttt ttggcgtttc gcgtcagtca atagaagttt 360
 cagatctaca ttaggaagaa ccagaaaata gccattaatg ctttcagcat agcacagcat 420
 agcagctgtg tatactttaa ataagatgta gactggtttg catttggaag ggttttgtgt 480
 aagaaaagca atacttgagg taaaacaaga gaaaaaaaaa cactttacta actaatatcc 540
 aatcctttat ttttttgcag aa atg aaa ttc tca act gcc gtt act acg ttg 592
 Met Lys Phe Ser Thr Ala Val Thr Thr Leu 10
 1 5

att agt tct ggt gcc atc gtg tct gct tta cca cac gtg gat gtt cac 640
 Ile Ser Ser Gly Ala Ile Val Ser Ala Leu Pro His Val Asp Val His 25
 15 20

caa gaa gat gcc cac caa cat aag agg gcc gtt gcg tac aaa tac gtt 688
 Gln Glu Asp Ala His Gln His Lys Arg Ala Val Ala Tyr Lys Tyr Val 40
 30 35

tac gaa act gtt gtt gtc gat tct gat ggc cac act gta act cct gct 736
 Tyr Glu Thr Val Val Val Asp Ser Asp Gly His Thr Val Thr Pro Ala 55
 45 50

gct tca gaa gtc gct act gct gct acc tct gct atc att aca aca tct 784
 Ala Ser Glu Val Ala Thr Ala Ala Thr Ser Ala Ile Ile Thr Thr Ser 70
 60 65

gtg ttg gct cca acc tcc tcc gca gcc gct ggg ata gcc gct tcc att 832
 Val Leu Ala Pro Thr Ser Ser Ala Ala Ala Gly Ile Ala Ala Ser Ile 90
 75 80 85

gct gtt tca tct gct gcc tta gcc aag aat gag aaa atc tct gat gcc 880
 Ala Val Ser Ser Ala Ala Leu Ala Lys Asn Glu Lys Ile Ser Asp Ala 105
 95 100

gct gca tct gcc act gcc tca aca tct caa ggg gca tcc tcc tcc tcc 928
 Ala Ala Ser Ala Thr Ala Ser Thr Ser Gln Gly Ala Ser Ser Ser Ser 120
 110 115

tcc tcc tcc tcg gca act tct acc cta gaa agc agc tct gtt tct tca 976
 Ser Ser Ser Ser Ala Thr Ser Thr Leu Glu Ser Ser Ser Val Ser Ser 135
 125 130

tct agt gaa gaa gct gct cca aca tct act gtc gtg tca act tct tcc 1024
 Ser Ser Glu Glu Ala Ala Pro Thr Ser Thr Val Val Ser Thr Ser Ser 150
 140 145

gca acc caa tct agt gct tct tct gcc act aaa tct agt act tct tcc 1072
 Ala Thr Gln Ser Ser Ala Ser Ser Ala Thr Lys Ser Ser Thr Ser Ser 170
 155 160 165

act tca cca tct act tct act tct act tcc act tct tct act tcc tct 1120
 Thr Ser Pro Ser Thr Ser Thr Ser Thr Ser Thr Ser Thr Ser Ser 185
 175 180

tcc tct tcc tcc tcc tcc tcc tct tct tct tct tct ggc agt ggt 1168
 Ser Ser Ser Ser Ser Ser Ser Ser Ser Ser Ser Ser Gly Ser Gly 200
 190 195

agt atc tac ggt gat ttg gcc gac ttt tca ggc cca agt gag aaa ttc 1216

105040-2329300

Ser Ile Tyr Gly Asp Leu Ala Asp Phe Ser Gly Pro Ser Glu Lys Phe
 205 210 215
 caa gac ggc act att cca tgt gac aaa ttc cca tct ggt caa ggt gtc 1264
 Gln Asp Gly Thr Ile Pro Cys Asp Lys Phe Pro Ser Gly Gln Gly Val
 220 225 230
 att tct att gac tgg att ggc gag ggt gga tgg tcc ggt gtg gaa aac 1312
 Ile Ser Ile Asp Trp Ile Gly Glu Gly Gly Trp Ser Gly Val Glu Asn
 235 240 245 250
 acc gac act tcc act ggc ggt tca tgc aag gag ggg tcc tac tgt tcc 1360
 Thr Asp Thr Ser Thr Gly Gly Ser Cys Lys Glu Gly Ser Tyr Cys Ser
 255 260 265
 tac tcc tgc caa cca ggt atg tct aag acc caa tgg cca tcc gat caa 1408
 Tyr Ser Cys Gln Pro Gly Met Ser Lys Thr Gln Trp Pro Ser Asp Gln
 270 275 280
 cca tct gac ggt aga tct gtc ggg ggt ttg ttg tgt aaa aat ggt tat 1456
 Pro Ser Asp Gly Arg Ser Val Gly Gly Leu Leu Cys Lys Asn Gly Tyr
 285 290 295
 ttg tac cgt tct aac act gac gcg gat tac tta tgt gaa tgg ggt gtc 1504
 Leu Tyr Arg Ser Asn Thr Asp Ala Asp Tyr Leu Cys Glu Trp Gly Val
 300 305 310
 gag gct gcc tat gtt gtt tct aaa cta agc aag ggt gtc gcc att tgc 1552
 Glu Ala Ala Tyr Val Val Ser Lys Leu Ser Lys Gly Val Ala Ile Cys
 315 320 325 330
 aga acc gac tac ccg ggc act gaa aac atg gtt atc cca acc tat gtt 1600
 Arg Thr Asp Tyr Pro Gly Thr Glu Asn Met Val Ile Pro Thr Tyr Val
 335 340 345
 gaa ggg ggt agc tct ttg cca ttg acc gtt gtt gac caa gat act tac 1648
 Glu Gly Gly Ser Ser Leu Pro Leu Thr Val Val Asp Gln Asp Thr Tyr
 350 355 360
 ttt act tgg gaa ggc aaa aag aca tct gct caa tac tac gtt aat aac 1696
 Phe Thr Trp Glu Gly Lys Lys Thr Ser Ala Gln Tyr Tyr Val Asn Asn
 365 370 375
 gcc ggc gtc tca gtt gaa gat ggg tgt atc tgg ggt act tct gga tct 1744
 Ala Gly Val Ser Val Glu Asp Gly Cys Ile Trp Gly Thr Ser Gly Ser
 380 385 390
 ggt att ggt aac tgg gca cca tta aac ttt ggt gct ggc tcc act ggt 1792
 Gly Ile Gly Asn Trp Ala Pro Leu Asn Phe Gly Ala Gly Ser Thr Gly
 395 400 405 410
 gga gtg aca tac tta tca ttg att cct aac cca aac aac agc gac gca 1840
 Gly Val Thr Tyr Leu Ser Leu Ile Pro Asn Pro Asn Asn Ser Asp Ala
 415 420 425
 ttg aac tac aac gtc aag ata gtt gct gct gat gat tca tcc aat gtc 1888
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<210> 14
<211> 475
<212> PRT
<213> Saccharomyces cerevisiae
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 Thr Glu Asn Met Val Ile Pro Thr Tyr Val Glu Gly Gly Ser Ser Leu
 340 345 350
 Pro Leu Thr Val Val Asp Gln Asp Thr Tyr Phe Thr Trp Glu Gly Lys
 355 360 365
 Lys Thr Ser Ala Gln Tyr Tyr Val Asn Asn Ala Gly Val Ser Val Glu
 370 375 380
 Asp Gly Cys Ile Trp Gly Thr Ser Gly Ser Gly Ile Gly Asn Trp Ala
 385 390 395 400
 Pro Leu Asn Phe Gly Ala Gly Ser Thr Gly Gly Val Thr Tyr Leu Ser
 405 410 415
 Leu Ile Pro Asn Pro Asn Asn Ser Asp Ala Leu Asn Tyr Asn Val Lys
 420 425 430
 Ile Val Ala Ala Asp Asp Ser Ser Asn Val Ile Gly Glu Cys Val Tyr
 435 440 445
 Glu Asn Gly Glu Phe Ser Gly Gly Ala Asp Gly Cys Thr Val Ser Val
 450 455 460
 Thr Ser Gly Lys Ala His Phe Val Leu Tyr Asn
 465 470 475

<210> 15
 <211> 145
 <212> PRT
 <213> *Saccharomyces cerevisiae*

<220>
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 <222> (1)...(145)
 <223> Xaa = Any Amino Acid

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 Xaa Gly Xaa Ser Xaa Pro Xaa Xaa Val Xaa Xaa Xaa Asp Xaa Tyr Xaa
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 Xaa Trp Xaa Gly Lys Lys Thr Ser Ala Gln Tyr Tyr Xaa Asn Asn Xaa
 35 40 45
 Gly Val Ser Xaa Glu Asp Gly Cys Ile Trp Gly Thr Xaa Gly Ser Xaa
 50 55 60
 Xaa Gly Asn Trp Ala Pro Xaa Xaa Xaa Gly Ala Xaa Xaa Thr Xaa Gly
 65 70 75 80
 Xaa Thr Tyr Leu Ser Xaa Ile Pro Asn Pro Asn Xaa Xaa Xaa Ala Xaa
 85 90 95
 Asn Xaa Asn Xaa Lys Ile Val Ala Xaa Asp Xaa Xaa Xaa Xaa Val Xaa
 100 105 110
 Gly Xaa Cys Xaa Tyr Glu Xaa Gly Xaa Xaa Gly Xaa Gly Xaa Asp
 115 120 125
 Gly Cys Thr Val Ser Val Xaa Ser Gly Xaa Ala Xaa Phe Val Xaa Tyr
 130 135 140
 Xaa
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<210> 16
 <211> 60

<212> PRT
 <213> *Saccharomyces cerevisiae*

<400> 16
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 1 5 10 15
 Lys Ile Val Ala Ala Asp Asp Ser Ser Thr Val Asn Gly Glu Cys Ile
 20 25 30
 Tyr Glu Asn Gly Ser Phe Ser Ser Gly Gly Ser Asp Gly Cys Thr Val
 35 40 45
 Ser Val Thr Ala Gly Lys Ala Lys Phe Val Leu Tyr
 50 55 60

<210> 17
 <211> 16
 <212> PRT
 <213> *Saccharomyces cerevisiae*

<400> 17
 Leu Ala Thr Asp Gln Phe Gly Cys Arg Phe Leu Gln Lys Lys Leu Glu
 1 5 10 15

<210> 18
 <211> 16
 <212> PRT
 <213> *Saccharomyces cerevisiae*

<400> 18
 Leu Ile Leu Asp Pro Phe Gly Asn Tyr Leu Val Asp Lys Ile Cys Asp
 1 5 10 15

<210> 19
 <211> 16
 <212> PRT
 <213> *Saccharomyces cerevisiae*

<400> 19
 Ile Ser Ile Asn Gln Tyr Gly Thr Arg Ser Leu Gln Lys Ile Ile Asp
 1 5 10 15

<210> 20
 <211> 15
 <212> PRT
 <213> *Saccharomyces cerevisiae*

<400> 20
 Leu Ile Asn Asp Ile Asn Gly His Val Ile Gln Lys Cys Ile Phe
 1 5 10 15

<210> 21
 <211> 16
 <212> PRT
 <213> *Saccharomyces cerevisiae*

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<210> 27
<211> 16
<212> PRT
<213> Saccharomyces cerevisiae
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<210> 33
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<212> PRT
<213> Drosophila
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<400> 33

<210> 34

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<210> 35

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<210> 39

<212> PRT

<213> Drosophila

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<210> 40
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<212> PRT
<213> Drosophila
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<210> 41
<211> 16
<212> PRT
<213> Homo sapiens
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<212> PRT
<213> Homo sapiens
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<212> PRT
<213> Homo sapiens
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<211> 16
<212> PRT
<213> Homo sapiens
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<210> 45
<211> 16
<212> PRT
<213> Homo sapiens
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<400> 45
Leu Ser Thr His Pro Tyr Gly Cys Arg Val Ile Gln Arg Ile Leu Gln
1 5 10 15

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<210> 46
<211> 16
<212> PRT
<213> Homo sapiens
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<400> 46
Leu Val Gln Asp Gln Tyr Gly Asn Tyr Val Ile Gln His Val Leu Glu
1 5 10 15

```
<210> 47
<211> 16
<212> PRT
<213> Homo sapiens
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```

<400> 47
Val Leu Ser Gln His Phe Ala Ser Asn Val Val Glu Lys Cys Val Thr
      1           5           10           15

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<210> 48
<211> 16
<212> PRT
<213> Homo sapiens
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<400> 48
Met Met Lys Asp Gln Tyr Ala Asn Tyr Val Val Gln Lys Met Ile Asp
1 5 10 15

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